



## UNITED STATES DEPARTMENT OF COMMERCE Patent and Trademark Office

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18N1/1105 -

DAVID A. KALOW LIEBERMAN & NOWAK 292 MADISON AVE., 8TH FLOOR NEW YORK, NY 10017

# NOTICE OF ALLOWANCE AND ISSUE FEE DUE

| ☐ Note attached communication from the Examiner                  |  |
|--|--|
| This notice is issued in view of applicant's communication filed |  |

| SERIES CODE/SERIAL NO.      | FILING DATE | TOTAL CLAIMS | EXAMINER AND GROUP ART | UNIT | DATE MAILED |
|-----------------------------|-------------|--------------|------------------------|------|-------------|
| 07/130,070                  | 12/08/87    | 042          | MARSCHEL, A            | 1807 | 11/05/93    |
| First Named Applicant WARD, |             | DAVI         | D C.                   |      | ····        |

TITLE OF INVENTION METHODS OF USING LABELED NUCLEOTIDES (AS AMENDED)

|   | ATTY'S DOCKET NO. | CLASS-SUBCLASS | BATCH NO. | APPLN. TYPE | SMALL ENTITY | FEE DUE   | DATE DUE |
|---|-------------------|----------------|-----------|-------------|--------------|-----------|----------|
| 1 | ENZ-1 (CONT) D    | 435-006.       | 000 S2    | 9 UTILI1    | ry no        | \$1170.00 | 02/07/94 |

THE FEE DUE IS THE AMOUNT IN EFFECT AT THIS TIME. IF THE AMOUNT OF THE ISSUE FEE INCREASES PRIOR TO PAYMENT, APPLICANT WILL BE NOTIFIED OF THE BALANCE OF ISSUE FEE DUE.

THE APPLICATION IDENTIFIED ABOVE HAS BEEN EXAMINED AND IS ALLOWED FOR ISSUANCE AS A PATENT.

### PROSECUTION ON THE MERITS IS CLOSED.

THE ISSUE FEE MUST BE PAID WITHIN <u>THREE MONTHS</u> FROM THE MAILING DATE OF THIS NOTICE OR THIS APPLICATION SHALL BE REGARDED AS ABANDONED. THIS STATUTORY PERIOD CANNOT BE EXTENDED.

### HOW TO RESPOND TO THIS NOTICE:

- I. Review the SMALL ENTITY Status shown above. If the SMALL ENTITY is shown as YES, verify your current SMALL ENTITY status:
- A. If the status is changed, pay twice the amount of the FEE DUE shown above and notify the patent and Trademark Office of the change in status, or
  - B. If the Status is the same, pay the FEE DUE shown above.

- If the SMALL ENTITY is shown as NO:
- A. Pay FEE DUE shown above, or
- B. File verified statement of Small Entity Status before, or with, pay of 1/2 the FEE DUE shown above.

- II. Part B of this notice should be completed and returned to the Patent and Trademark Office (PTO) with your ISSUE FEE. Even if the ISSUE FEE has already been paid by charge to deposit account, Part B should be completed and returned. If you are charging the ISSUE FEE to your deposit account, Part C of this notice should also be completed and returned.
- III. All communications regarding this application must give series code (or filing date) and serial number. Please direct all communications prior to issuance to Box ISSUE FEE unless advised to contrary.

IMPORTANT REMINDER: Patents issuing on applications filed on or after Dec. 12, 1980 may require payment of maintenance fees. It is patentee's responsibility to ensure timely payment of maintenance fees when due.



#### UNITED STATES DEPARTMENT OF COMMERCE **Patent and Trademark Office**

Address: COMMISSIONER OF PATENTS AND TRADEMARKS Washington, D.C. 20231

SERIAL NUMBER **FILING DATE** FIRST NAMED APPLICANT ATTORNEY DOCKET NO. 07/130,070 12/08/87 WARD EXAMNER 1 (CONT) D MARSCHEL, A 18N1/1105 ART UNIT PAPER NUMBER DAVID A. KALOW LIEBERMAN & NOWAK 292 MADISON AVE., 8TH FLOOR NEW YORK, NY 10017 DATE MAILED?

11/05/93

#### **NOTICE OF ALLOWABILITY**

| PAR  | TI         | 1 ATTI 1 12 7000 AT 1 DAL 110 7 80 0 0 0 1 1 1 110-7-9  |
|------|------------|---|
| 1.   | 29         | This communication is responsive to Andt I filed 8-27-93, term. disol, filed 10-7-93, and Suppl. declar, Ital 10-7-9.   |
| 2.   | ×          | All the claims being allowable, PROSECUTION ON THE MERITS IS (OR REMAINS) CLOSED in this application. If not included herewith (or previously mailed), a Notice Of Allowance And Issue Fee Due or other appropriate communication will be sent in due   |
| 3.   | Ø          | The allowed claims are 126-130, 134-136, 142, 143, 150, 151-154, 156, 157, 159-183, +185; renumbered as 33-42 and 1-32, respectively.   |
| 4.   |            | The drawings filed on are acceptable.   |
|      |            | Acknowledgment is made of the claim for priority under 35 U.S.C. 119. The certified copy has [_] been received. [_] not been received. [_] been filed in parent application Serial No   |
| 6.   | X          | Note the attached Examiner's Amendment.   |
| 7.   | Ø          | Note the attached Examiner Interview Summary Record, PTOL-413.  |
| 8.   | Ø          | Note the attached Examiner's Statement of Reasons for Allowance.  |
| 9.   |            | Note the attached NOTICE OF REFERENCES CITED, PTO-892.  |
|      |            | Note the attached INFORMATION DISCLOSURE CITATION, PTO-1449.  |
|      |            |   |
| PAR  |            |   |
| FRC  | M          | RTENED STATUTORY PERIOD FOR RESPONSE to comply with the requirements noted below is set to EXPIRE THREE MONTHS  THE "DATE MAILED" indicated on this form. Failure to timely comply will result in the ABANDONMENT of this application.  ions of time may be obtained under the provisions of 37 CFR 1.136(a). |
| LAIC | ,,,,       | ons of time may be obtained under the provisions of 37 Gr N 1. Iso(a).  |
| 1. [ |            | Note the attached EXAMINER'S AMENDMENT or NOTICE OF INFORMAL APPLICATION, PTO-152, which discloses that the oath or declaration is deficient. A SUBSTITUTE OATH OR DECLARATION IS REQUIRED.   |
| 2. [ |            | APPLICANT MUST MAKE THE DRAWING CHANGES INDICATED BELOW IN THE MANNER SET FORTH ON THE REVERSE SIDE<br>OF THIS PAPER.   |
| а    | . <b>C</b> | ☐ Drawing informalities are indicated on the NOTICE RE PATENT DRAWINGS, PTO-948, attached hereto or to Paper No CORRECTION IS REQUIRED.   |
| b    | ). [       | ☐ The proposed drawing correction filed on has been approved by the examiner. CORRECTION IS REQUIRED.   |
| С    | . [        | Approved drawing corrections are described by the examiner in the attached EXAMINER'S AMENDMENT. CORRECTION IS  |
| á    | í. C       | ☐ Formal drawings are now REQUIRED.   |
|      |            |   |
| A    |            | species to this latter should include in the unreal right hand source the following information from the NOTICE OF ALL OWANGE   |
|      |            | sponse to this letter should include in the upper right hand corner, the following information from the NOTICE OF ALLOWANCE<br>SUE FEE DUE: ISSUE BATCH NUMBER, DATE OF THE NOTICE OF ALLOWANCE, AND SERIAL NUMBER.   |
| Atta | chn        | nents:  |
|      |            | iner's Amendment _ Notice of Informal Application, PTO-152  |
|      | _          | iner Interview Summary Record, PTOL- 413 Notice re Patent Drawings, PTO-948   |
|      |            | ons for Allowance Listing of Bonded Draftsmen   |
|      |            | e of References Cited, PTO-892 Other  |
| _ In | torr       | nation Disclosure Citation, PTO-1449  |

MARGARET PARR SUPERVISORY PATENT EXAMINER **GROUP 1800** 

TOL-37 (REV. 4-89) \*

Applicants are hereby informed that the amendment filed 8/27/93, which was previously indicated as denied entry in the Advisory Action mailed 9/24/93, has been entered.

An Examiner's Amendment to the record appears below. the changes and/or additions be unacceptable to applicant, an amendment may be filed as provided by 37 C.F.R. § 1.312. ensure consideration of such an amendment, it MUST be submitted no later than the payment of the Issue Fee.

An extension of time under 37 C.F.R. § 1.136(a) is required in order to make an Examiner's Amendment which places this application in condition for allowance. During a telephone conversation conducted on 10/25/93, Ron Fedus requested an extension of time for an additional 2 months and authorized the Commissioner to charge Deposit Account No. 05-1135 the required fee for this extension and authorized the following Examiner's Should the changes and/or additions be unacceptable Amendment. to applicant, an amendment may be filed as provided by 37 C.F.R. § 1.312. To ensure consideration of such an amendment, it MUST be submitted no later than the payment of the Issue Fee.

## In the claims:

In claim 150, line 15, after "wherein B and A are"; insert --covalently--.

Cancel claim 155, without prejudice.

In claim 157, line 3, after "enzyme"; insert --capable of depositing insoluble reaction products--.

Cancel claim 158, without prejudice.

In claim 161, lines 3-4, delete "reacting with a substrate to form a detectable product"; and insert therefor --depositing insoluble reaction products--.

In claim 168, line 2, delete "compound"; and insert therefor -- of said compounds -- .

In claim 168, line 3, delete "compound"; and insert therefor -- of said compounds -- .

In claim 171, lines 3-4, delete "reacting with a substrate to form a detectable product"; and insert therefor --depositing insoluble reaction products--.

The following is an Examiner's Statement of Reasons for Allowance:

The invention is directed to methods of detection of nucleic acids using a hybridization probe that contains modified bases that are modified by covalent labeling with an A moiety attached at either the 7-position of 7-deazapurine or the 5-position of pyrimidine. The A moiety contains at least three carbon atoms and mediates the production of a detectable signal.

The prior art of record teaches various nucleotide base modifications including at the above two linkage points but this prior art does not teach or suggest that such modified bases can be incorporated into nucleic acid probes by enzymatic means or that probes containing such modifications are still efficient hybridization probes. It is noted that a divisional application to the instant application has been previously issued as patent number 4,711,955; directed to biotin and iminobiotin label types

as species of the above discussed A moiety.

The terminal disclaimer has been instantly filed on 10/27/93 to obviate and prevent an obviousness-double patenting rejection over claims 15-20 of patent number 4,711,955 which suggests hybridization methods via the hybrid formation cited in said claims 15-20.

1807

The supplemental Declaration under 37 CFR § 1.131 has been instantly filed on 10/27/93 to perfect a previous Declaration and to obviate and prevent a rejection over the reference cited as Langer III; Abstract 1153, J. Supramolecular Structure and Cellular Biochemistry (Suppl. 5) (1981) of a presentation given on December 11, 1980.

Any comments considered necessary by applicant must be submitted no later than the payment of the Issue Fee and, to avoid processing delays, should preferably accompany the Issue Fee. Such submissions should be clearly labeled "Comments on Statement of Reasons for Allowance."

this application should be directed to the Group receptionist

Any inquiry of a general nature or relating to the status of

whose telephone number is (703) 308-0196.  $\Lambda M$ 

A. MARSCHEL:am

October 26, 1993

MARGARET PARR SUPERVISORY PATENT EXAMINER GROUP 1800